It would have been obvious to one of ordinary skill in the art at the time the invention was made to treat stroke patients employing a combination of phosphodiesterase inhibitors and training.

One of ordinary skill in the art would have been motivated to treat stroke patients employing a combination of phosphodiesterase inhibitors and training because both PDE inhibitors and cognitive training are known to be useful in method of treating stroke. One of ordinary skill in the art would have reasonably expected a combination of training and PDE inhibitors to be useful in treating stroke patients.

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Mojdeh Bahar whose telephone number is (703) 305-1007. The examiner can normally be reached on (703) 305-1007 from 8:30 a.m. to 6:30 p.m. Monday, Tuesday, Thursday and Friday.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Sreeni Padmanabhan can be reached on (703) 305-1877. The fax phone number for the organization where this application or proceeding is assigned is (703) 308-4556.

Any inquiry of a general nature or relating to the status of this application or proceeding should be directed to the receptionist whose telephone number is (703) 308-1235.

Mojdeh Bahar
Patent Examiner
April 29, 2003

SREENI PADMANABHAN
PRIMARY EXAMINER 5/2/03

PTO-1449 REPRODUCED

SUPPLEMENTAL INFORMATION DISCLOSURE CITATION IN AN APPLICATION

August 29, 2002 (Use several sheets if necessary)

ATTORNEY DOCKET NO.	APPLICATION NO.
1314.2004-001	09/927,914
APPLICANT: Timothy P. Tully *et al.*	
FILING DATE: August 10, 2001	GROUP: 1617

U.S. PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	ISSUE DATE / PUBLICATION DATE	NAME

FOREIGN PATENT DOCUMENTS

		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION YES	TRANSLATION NO

OTHER DOCUMENTS ***(Including Author, Title, Date, Pertinent Pages, Etc.)***

MW	AV5	Milner, B. *et al.*, "Cognitive Neuroscience and the Study of Memory," *Neuron*, 20(3):445-468 (1998).
MW	AW5	Bevilaqua, L. *et al.*, "Drugs Acting Upon the Cyclic Adenosine Monophosphate/Protein Kinase A Signalling Pathway Modulate Memory Consolidation When Given Late After Training into Rat Hippocampus but not Amygdala," *Behavioural Pharmacology*, 8(4):331-338 (1997).

EXAMINER: MH

DATE CONSIDERED: 04/29/03

PTO-1449 REPRODUCED	ATTORNEY DOCKET NO. 1314.2004-001	APPLICATION NO. 09/927,914
INFORMATION DISCLOSURE CITATION IN AN APPLICATION February 2, 2002 (Use several sheets if necessary)	APPLICANT Timothy P. Tully and Filippo Cavalier	
	FILING DATE August 10, 2001	GROUP 1619

OTHER DOCUMENTS ***(Including Author, Title, Date, Pertinent Pages, Etc.)***

	AW3	Silva, A.J. *et al.*, "CREB and Memory," *Annu. Rev. Neurosci.*, 21:127-148 (1998).
	AX3	Bourtchouladze, R. *et al*, "Different Training Procedures Recruit Either One or Two Critical Periods for Contextual Memory Consolidation, Each of Which Requires Protein Synthesis and PKA," *Learning & Memory*, 5:365-374 (1998).
	AY3	Yin, J.C.P. *et al.*, "Induction of a Dominant Negative CREB Transgene Specifically Blocks Long-Term Memory in Drosophila," *Cell*, 79:49-58 (1994).
	AZ3	Yin, J.C.P. *et al.*, "CREB as a Memory Modulator: Induced Expression of a dCREB2 Activator Isoform Enhances Long-Term Memory in Drosophila," *Cell*, 81:107-115 (1995).
	AR4	Josselyn, S.A. *et al.*, "Overexpression of CREB in the Amygdala Facilitates the Formation of Long-Term Memory Measured with Fear Potentiated Startle in Rats," *Society for Neuroscience*, Vol. 24, Abstract 365.10 (1998).
	AS4	Kogan, J.H. *et al.*, "Spaced Training Induces Normal Long-Term Memory in CREB Mutant Mice," *Current Biology*, 7:1-11 (1996).
	AT4	Bartsch, D. *et al.*, "Aplysia CREB2 Represses Long-Term Facilitation: Relief of Repression Converts Transient Facilitation into Long-Term Functional and Structural Change," *Cell*, 83:979-992 (1995).
	AU4	Pedreira, M.E., "Massed and Spaced Training Build Up Different Components of Long-Term Habituation in the Crab *Chasmagnathus*," *Animal Learning & Behavior*, 26(3):34-43 (1998).
	AV4	Bourtchuladze, R. *et al.*, "Deficient Long-Term Memory in Mice with a Targeted Mutation of the cAMP-Responsive Element-Binding Protein," *Cell*, 79:59-68 (1994).
	AW4	Tully, T. *et al.*, "Genetic Dissection of Consolidated Memory in Drosophila," *Cell*, 79:35-47 (1994).
	AX4	Guzowski, J.F. and McGaugh, J.L., "Antisense Oligodeoxynucleotide-Mediated Disruption of Hippocampal cAMP Response Element Binding Protein Levels Impairs Consolidation of Memory for Water Maze Training," *Proc. Natl. Acad. Sci. USA*, 94:2693-2698 (1997).
	AY4	Lamprecht, R. *et al.*, "cAMP Response Element-Binding Protein in the Amygdala is Required for Long-but not Short-Term Conditioned Taste Aversion Memory," *The Journal of Neuroscience*, 17(21):8443-8450 (1997).
	AZ4	Impey, S. *et al.*, "Stimulation of cAMP Response Element (CRE)-Mediated Transcription During Contextual Learning," *Nature Neuroscience*, 1(7):595-601 (1998).

EXAMINER	DATE CONSIDERED
	04/29/03

PTO-1449 REPRODUCED	ATTORNEY DOCKET NO. 1314.2004-001	APPLICATION NO. 09/927,914
INFORMATION DISCLOSURE CITATION IN AN APPLICATION	APPLICANT Timothy P. Tully and Filippo Cavallaro	
February 2, 2002 (Use several sheets if necessary)	FILING DATE August 10, 2001	GROUP 1619

OTHER DOCUMENTS (*Including Author, Title, Date, Pertinent Pages, Etc.*)

	AR5	Moore, A.N. *et al.*, "Neuronal Activity Increases the Phosphorylation of the Transcription Factor cAMP Repsonse Element-binding Protein (CREB) in Rat Hippocampus and Cortex," *The Journal of Biological Chemistry*, 271(24):14214-14220 (1996).
	AS5	Murphy, D.D. and Segal, M., "Morphological plasticity of Dendritic Spines in Central Neurons is Mediated by Activation of cAMP Response Element Binding Protein," *Proc. Natl. Acad. Sci. USA*, 94:1482-1487 (1997).
	AT5	Liu, F.C. and Graybiel, A.M., "Spatiotemporal Dynamics of CREB Phosphorylation: Transient Versus Sustained Phosphorylation in the Developing Striatum," *Neuron*, 17:1133-1144 (1996).
	AU5	Dubnau, J. and Tully, T., "Gene Discovery in Drosophila: New Insights for Learning and Memory," *Annu. Rev. Neurosci.*, 21:407-444 (1998).

EXAMINER	DATE CONSIDERED 04/29/03

PTO-1449 REPRODUCED

INFORMATION DISCLOSURE CITATION IN AN APPLICATION

February 2, 2002

(Use several sheets if necessary)

ATTORNEY DOCKET NO.	APPLICATION NO.
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APPLICANT: Timothy P. Tully and Filippo Cavalie	
FILING DATE: August 10, 2001	GROUP: 1619

U.S. PATENT DOCUMENTS

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FOREIGN PATENT DOCUMENTS

		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION YES NO

OTHER DOCUMENTS ***(Including Author, Title, Date, Pertinent Pages, Etc.)***

MS	AU	Ikezu, T. *et al.*, "Negative Transactivation of cAMP Response Element by Familial Alzheimer's Mutants of APP," *The EMBO Journal*, 15(10):2468-2475 (1996).
	AV	Sato, N. *et al.*, "Elevated Amyloid β Protein (1-40) Level Induces CREB Phosphorylation at Serine-133 via p44/42 MAP Kinase (Erk1/2)-Dependent Pathway in Rat Pheochromocytoma PC12 Cells," *Biochemical and Biophysical Research Communications*, 232:637-642 (1997).
	AW	Yamamoto-Sasaki, M. *et al.*, "Impaired Phosphorylation of Cyclic AMP Response Element Binding Protein in the Hippocampus of Dementia of the Alzheimer Type," *Brain Research*, 824:300-303 (1999).
	AX	Blendy, J.A., "Effects of Kainic Acid Induced Seizures on Immediate Early Gene Expression in Mice with a Targeted Mutation of the CREB Gene," Brain Research, 681:8-14 (1995).
	AY	Tanaka, K. *et al.*, "Temporal Profile of CREB Phosphorylation After Focal Ischemia in Rat Brain," *NeuroReport*, 10:2245-2250 (1999).
	AZ	Young, D. *et al.*, "Environmental Enrichment Inhibits Spontaneous Apoptosis, Prevents Seizures and is Neuroprotective," *Nature Medicine*, 5(4):448-453 (1999).
MS	AR2	Pandey, S.C. *et al.*, "Involvement of the Cyclic AMP-Responsive Element Binding Protein Gene Transcription Factor in Genetic Preference for Alcohol Drinking Behavior," *Alcohol. Clin. Exp. Res.*, 23(9):1425-1434 (1999).

EXAMINER	DATE CONSIDERED
MS	04/29/03

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INFORMATION DISCLOSURE CITATION IN AN APPLICATION

February 2, 2002

(Use several sheets if necessary)

ATTORNEY DOCKET NO.	APPLICATION NO.
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APPLICANT: Timothy P. Tully and Filippo Cavanna	
FILING DATE: August 10, 2001	GROUP: 1619

U.S. PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB-CLASS	FILING DATE IF APPROPRIATE
MP	AA	5,929,223	27-JULY-1999	Tully *et al.*	536	23.5	
MP	AB	6,051,559	18-APR-2000	Tully et al.	514	44	
	AC						
	AD						
	AE						
	AF						
	AG						
	AH						
	AI						
	AJ						
	AK						

FOREIGN PATENT DOCUMENTS

		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION YES NO
MP	AL	WO 96/11270	18-APR-1996	PCT			
	AM						
	AN						
	AO						
	AP						
	AQ						

OTHER DOCUMENTS ***(Including Author, Title, Date, Pertinent Pages, Etc.)***

MP	AR	Ahn, S. *et al.*, "A Late Phase of Cerebellar Long-Term Depression Requires Activation of CaMKIV and CREB," *Neuron*, 23:559-568 (1999).
MP	AS	Pham, T.A. *et al.*, "CRE-Mediated Gene Transcription in Neocortical Neuronal Plasticity during the Developmental Critical Period," *Neuron*, 22:63-72 (1999).
MP	AT	Glazewski, S. *et al.*, "Impaired Experience-Dependent Plasticity in Barrel Cortex of Mice Lacking the Alpha and Delta Isoforms of CREB," *Cerebral Cortex*, 9:249-256 (1999).

EXAMINER [signature]

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